SEARCH REQUEST FORM

Scientific and Technical Information Center

Art Unit:	Phone Number 30	Examiner # : Serial Number:	
fail-Box and Bldg/Room	Location:	Results Format Preferred (circ	le): PAPER DISK E-MAI
more than one search	1 is submitted, please p *********	orioritize searches in order of	need. *****************
nclude the elected species or stility of the invention. Define	structures, keywords, synonyr	describe as specifically as possible the sins, acronyms, and registry numbers, an pecial meaning. Give examples or relevints, and abstract.	d:combine with the concept or
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For Sequence Searches Only*	Please include all pertinent info	rmation (parent, child, divisional, or issue	d patent numbers) alone with the
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TAFF USE ONLY	Type of Search	Vendors and cost	where applicable
ircher:	NA Sequence (#)	STN	
rcher Phone #:	AA Sequence (#)	Dialog	
rcher Location:	Structure (#)	Questel/Orbit	
e Searcher Picked Up: 10	10/02 Bibliographic	Dr.Link	•
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te Completed: 10	ル (のV Litigation	Lexis/Nexis	·

PTO-1590 (8-01)

Clerical Prep Time:

Online Time:

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Patent Family

Manual of Patent Examining Pr cedure, S ction 713:04 Substanc of Interview must Be Mad of R cord

A complete written statement as to the substance of any face-to-face or telephone interview with regard to an application must be made of record in the application, whether or not an agreement with the examiner was reached at the interview.

§1.133 Interviews

(b) In every instance where reconsideration is requested in view of an interview with an examiner, a complete written statement of the reasons presented at the interview as warranting favorable action must be filed by the applicant. An interview does not remove the necessity for response to Office action as specified in §§ 1.111,1.135. (35 U.S.C.132)

1.2. Business to be transacted in writing. All business with the Patent or Trademark Office should be transacted in writing. The personal attendance of applicants or their attorneys or agents at the Patent and Trademark Office is unnecessary. The action of the Patent and Trademark Office will be based exclusively. on the written record in the Office. No attention will be paid to any alleged oral promise, stipulation, or understanding in relation to which there is disagreement or

The action of the Patent and Trademark Office cannot be based exclusively on the written record in the Office if that record is itself incomplete through the failure to record the substance of interviews.

It is the responsibility of the applicant or the attorney or agent to make the substance of an interview of record in the application file, unless the examiner indicates he or she will do so. It is the examiner's responsibility to see that such a record is made and to correct material inaccuracies which bear directly on the question of patentability.

Examiners must complete a two-sheet carbon interleaf Interview Summary Form for each interview held after January 1, 1978 where a matter of substance has been discussed during the interview by checking the appropriate boxes and filling in the blanks in neat handwritten form using a ball-point pen. Discussions regarding only procedural matters, directed solely to restriction requirements for which interview recordation is otherwise provided for in Section 812.01 of the Manual of Patent only procedural matters, directed solely to restriction requirements for which interview recordation is otherwise procedure, or pointing out typographical errors or unreadable script in Office actions or the like, are excluded from the interview-recordation procedures below.

The Interview Summary Form shall be given an appropriate paper number; placed in the right than portion of the file, and listed on the "Contents" list on the file wrapper. The docket and serial register cards need not be updated to reflect interviews. In a personal interview, the duplicate copy of the Form is removed and given to the applicant (or attorney or agent) at the conclusion of the interview. In the case of a telephonic interview, the copy is mailed to the applicant's correspondence address either with or prior to the next official communication. If additional correspondence from the examiner is not likely before an allowance or if other circumstances dictate, the Form should be maited promptly after the telephonic interview rather than with the next official communication.

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Ad boy not 19 year reached 19 was not resulted.

Earlink and who demonstration consumed. If her little is you, ories described

The Form provides for recordation of the following information:

- -Serial Number of the application
- -Name of applicant
- -Name of examiner
- -Date of interview
- -Type of interview (personal or telephonic)
- Name of participant(s)) (applicant, attorney or agent, etc.)
- -An indication whether or not an exhibit was shown or a demonstration conducted

Fregue and Line of the

- -An identification of the claims discussed
- -An identification of the specific prior art discussed......
- -An indication whether an agreement was reached and if so, a description of the general nature of the agreement (may be by attachment of a copy of amendments or claims agreed as being allowable). (Agreements as to allowability are tentative and do not restrict further action by the examiner to the
- The signature of the examiner who conducted the interview
- -Names of other Patent and Trademark Office personnel present.

The Form also contains a statement reminding the applicant of his responsibility to record the substance of the interview.

It is deslicable that the examiner or ally committee applicant of his obligation to record the substance of the interview in each case unless both applicant and examiner agree that the examiner will record same. Where the examiner agree that the examiner will record same. Where the examiner agree that the interview or when it is adequately recorded on the Form or in an attachment to the Form, the examiner should checke box at the bottom of the Form informing the applicant that he need not supplement the Form by submitting a separate record of the substance of the interview.

It should be noted, however, that the Interview Summary Forth with not normally be considered a complete and proper recordation of the interview unless it includes, or is supplemented by the applicant or the examiner to include all on the applicable items required below concerning the substance of the interview?

A complete and proper recordation of the substance of any interview should include at least the following applicable items:

- -1) A brief description of the nature of any exhibit shown or any demonstration conducted;
- 2) an identification of the claims discussed,
- 3) antidentification of specific prior artidiscussed. In the least of the first account of the first prior articiscussed and the first account of the first
- 4) antidentification of the principal proposed amendments of a substantive nature discussed, unless these are already described on the interview Summary. 373. Form completed by the examiner,
- 5) a brief identification of the general thrust of the principal arguments presented to the examiner. The identification of arguments need not be lengthy or elaborate. A verbatim or highly detailed description of the arguments is not required. The identification of the arguments is sufficient if the general nature. or thrust of the principal arguments made to the examiner can be understood in the context of the application file. Of course, the applicant may desire to or thrust of the principal arguments made to the examiner can be understood in the context of the examiner can be understood in the context of the examiner can be understood in the context of the examiner can be understood in the context of the examiner can be understood in the context of the examiner can be understood in the examiner can be understood in the context of the examiner can be understood in the context of the examiner can be understood in the examiner can be unders
- 6) a general indication of any other pertinent matters discussed, and 30 miles a linearly described in the Interview Stimmary Form completed by the examiner.

Examiners are expected to carefully review the applicant's record of the substance of an interview. If the record is not complete or accurate, the examiner will give the applicant one month from the date of the notifying letter or the remainder of any period for response, whichever is longer, to complete the response and thereby avoid abandonment of the application (37 CFR 1.135(c)).

Examiner to Check for Accuracy

combined into a world fixed higher actinomials Applicant's summary of what took place at the interview should be carefully checked to determine the accuracy of any argument or statement attributed to the examiner during the interview. If there is aminaccuracy and it bears directly on the question of patentability, it should be pointed out in the next Office letter; if the claims are allowable for other reasons of record, the examiner should send a letter setting forth his or her version of the statement attributed to him. If the record is complete and accurate, the examiner should place the indication "interview record OK" on the paper recording the substance of the interview along with the date and the examiner's initials.

> CORRAINE SPECTOR PAIMARY EXAMINER

```
; Sequence 3, Application US/09813398
; GENERAL INFORMATION:
  APPLICANT: Bruce D. Weintraub
  APPLICANT: Mariusz W. Szkudlinski
; APPLICANT: University of Maryland
  TITLE OF INVENTION: CYSTINE KNOT GROWTH FACTOR MUTANTS
; FILE REFERENCE: UOFMD.003C1
  CURRENT APPLICATION NUMBER: US/09/813,398
  CURRENT FILING DATE: 2001-03-20
  PRIOR APPLICATION NUMBER: PCT/US99/05908
  PRIOR FILING DATE: 1999-03-19
  PRIOR APPLICATION NUMBER: PCT/US98/19772
  PRIOR FILING DATE: 1998-09-22
  NUMBER OF SEQ ID NOS: 41
  SOFTWARE: FastSEQ for Windows Version 4.0
; SEQ ID NO 3
   LENGTH: 141
   TYPE: PRT
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US-09-813-398-3
PSKEPLRPRCRPINATLAVEKEGCPVCITVNTTICAGYCPTMTRVLQGVLPALPQVVCNYRDVRFESIRL
PGCPRGVNPVVSYAVALSCQCALCRRSTTDCGGPKDHPLTCDDPRFQDSSSSKAPPPSLPSPSRLPGPSD
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Property values tagged with IC are from the ZIC/VINITI data file jan.delaval@uspto.gov provided by InfoChem.

STRUCTURE FILE UPDATES: 10 OCT 2002 HIGHEST RN 460706-73-4 DICTIONARY FILE UPDATES: 10 OCT 2002 HIGHEST RN 460706-73-4

TSCA INFORMATION NOW CURRENT THROUGH MAY 20, 2002

Please note that search-term pricing does apply when conducting SmartSELECT searches.

Crossover limits have been increased. See HELP CROSSOVER for details.

Experimental and calculated property data are now available. See HELP PROPERTIES for more information. See STNote 27, Searching Properties in the CAS Registry File, for complete details: http://www.cas.org/ONLINE/STN/STNOTES/stnotes27.pdf

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L19 O SEA FILE=REGISTRY ABB=ON PLU=ON PSKEPLRPRCRPINATLAVEKEGCPVCIT VNTTICAGYCPTMTRVLQGVLPALPQVVCNYRDVRFESIRLPGCPRGVNPVVSYAVALSCOCA LCRRSTTDCGGPKDHPLTCDDPRFQDSSSSKAPPPSLPSPSRLPGPSDT/SQSP

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L21

.....YCPTMTRVLQGVLPALPQVV......SCQCA

LCRRSTTDCGGPKDHPLTCDDPRFQDSSSSKAPPPSLPSPSRLPGPSDT/SQSP

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L21 10 SYCPTMTRVLQGVLPALPQVV.... SAV L21 SPECTOR813/A

FILE 'HCAPLUS' ENTERED AT 17:05:20 ON 11 OCT 2002

1.22 16 S L21

O S L22 AND (WEINTRAUB B? OR SZKUDLINSKI M?)/AU L23

L24 1 S WO99-US5908/AP, PRN

L25 0 S L22 AND L24

L26 11 S L22 AND (PY<=1997 OR PRY<=1997 OR AY<=1997)

SEL HIT RN

FILE 'REGISTRY' ENTERED AT 17:08:32 ON 11 OCT 2002

L27 7 S E1-E7

7 S L27 AND L21 L28

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L28 ANSWER 1 OF 7 REGISTRY COPYRIGHT 2002 ACS

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RN
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CN
    36: PN: US6238890 SEQID: 36 unclaimed protein (9CI) (CA INDEX NAME)
    PROTEIN SEQUENCE
FS
SQL 181
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Sequence | Patent
Source
      Reference
____+__+__
Not Given|US6238890
       |unclaimed
       |SEQID 36
SEO
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                         51 TTICAGYCPT MTRVLOGVLP ALPOVVCNYR DVRFESIRLP GCPRGVNPVV
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         151 PSRLPGPSDT PILPQGSGSG SGSAPDVQDC P
         ========
HITS AT:
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CI
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SR
    CA
LC
    STN Files:
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            1 REFERENCES IN FILE CA (1962 TO DATE)
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REFERENCE
          1: 135:1276
L28 ANSWER 2 OF 7 REGISTRY COPYRIGHT 2002 ACS
RN
    342058-80-4 REGISTRY
CN
    1-145-Gonadotropin, chorionic deriv. (human subunit .beta.) fusion protein
    with peptide fusion protein with 1-92-chorionic gonadotropin deriv. (human
    subunit .alpha.) (9CI) (CA INDEX NAME)
OTHER NAMES:
    39: PN: US6238890 SEQID: 39 claimed protein
CN
FS
    PROTEIN SEQUENCE
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Source | Reference
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         151 PSRLPGPSDT PILPQGSGSG SGSAPDVQDC PECTLQENPF FSQPGAPILQ
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     251 QHTACHCSTC YYHKS
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MF

Unspecified

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CI
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    CA
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             1 REFERENCES IN FILE CAPLUS (1962 TO DATE)
REFERENCE
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L28 ANSWER 3 OF 7 REGISTRY COPYRIGHT 2002 ACS
RN
    342058-60-0 REGISTRY
CN
    1-145-Gonadotropin, chorionic (human subunit .beta.) fusion protein with
    peptide fusion protein with 1-92-chorionic gonadotropin (human subunit
    .alpha.) (9CI) (CA INDEX NAME)
OTHER NAMES:
    3: PN: US6238890 SEQID: 3 claimed protein
CN
FS
    PROTEIN SEQUENCE
SQL 265
PATENT ANNOTATIONS (PNTE):
Sequence | Patent
Source | Reference
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        |claimed
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        1 MEMFQGLLLL LLLSMGGTWA SKEPLRPRCR PINATLAVEK EGCPVCITVN
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          101 SYAVALSCQC ALCRRSTTDC GGPKDHPLTC DDPRFQDSSS SKAPPPSLPS
          151 PSRLPGPSDT PILPQGSGSG SGSAPDVQDC PECTLQENPF FSQPGAPILQ
          _____
      201 CMGCCFSRAY PTPLRSKKTM LVQKNVTSES TCCVAKSYNR VTVMGGFKVE
      251 NHTACHCSTC YYHKS
HITS AT:
          20-160
MF
    Unspecified
CI
    MAN
SR
LC
    STN Files: CA, CAPLUS, USPATFULL
             1 REFERENCES IN FILE CA (1962 TO DATE)
             1 REFERENCES IN FILE CAPLUS (1962 TO DATE)
          1: 135:1276
REFERENCE
L28 ANSWER 4 OF 7 REGISTRY COPYRIGHT 2002 ACS
RN
    202016-40-8 REGISTRY
CN
    Gonadotropin, chorionic (human .beta.-subunit precursor) (9CI) (CA INDEX
    NAME)
OTHER NAMES:
    2: PN: US6319504 SEQID: 2 unclaimed protein
CN
CN
    Chorionic gonadotropin (human .beta.-subuint precursor)
CN
    Chorionic gonadotropin (human .beta.-subunit precursor)
FS
    PROTEIN SEQUENCE
SQL 165
PATENT ANNOTATIONS (PNTE):
Sequence | Patent
Source | Reference
======+==========
Not Given|US6319504
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|unclaimed |SEQID 2

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       1 MEMFQGLLLL LLLSMGGTWA SKEPLRPRCR PINATLAVEK EGCPVCITVN
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         101 SYAVALSCQC ALCRRSTTDC GGPKDHPLTC DDPRFQDSSS SKAPPPSLPS
         151 PSRLPGPSDT PILPQ
         _____
HITS AT:
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MF
    Unspecified
CI
    MAN
SR
    CA
    STN Files: CA, CAPLUS, TOXCENTER, USPATFULL
LC
            6 REFERENCES IN FILE CA (1962 TO DATE)
            3 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
            6 REFERENCES IN FILE CAPLUS (1962 TO DATE)
REFERENCE
          1: 136:1103
REFERENCE
          2: 134:4040
REFERENCE
          3:
            133:134164
REFERENCE
          4: 131:295923
REFERENCE
          5:
            128:124125
REFERENCE
          6: 128:124124
L28 ANSWER 5 OF 7 REGISTRY COPYRIGHT 2002 ACS
    195460-74-3 REGISTRY
RN
CN
    20-190-Tumor necrosis factor receptor p55 (human clone
    D.alpha.-TBP190hCG.beta.) fusion protein with peptide (synthetic linker)
    fusion protein with chorionic gonadotropin (human .beta.-subunit fragment)
    (9CI) (CA INDEX NAME)
OTHER NAMES:
CN
    8: PN: US6194177 SEQID: 8 claimed protein
FS
    PROTEIN SEQUENCE
SQL 336
PATENT ANNOTATIONS (PNTE):
Sequence | Patent
Source | Reference
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Not Given|US6194177
       Iclaimed
       |SEQID 8
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       1 SRTSLLLAFG LLCLPWLQEG SADSVCPQGK YIHPQNNSIC CTKCHKGTYL
      51 YNDCPGPGQD TDCRECESGS FTASENHLRH CLSCSKCRKE MGQVEISSCT
     101 VDRDTVCGCR KNQYRHYWSE NLFQCFNCSL CLNGTVHLSC QEKQNTVCTC
     151 HAGFFLRENE CVSCSNCKKS LECTKLCLPQ IENVKGTEDS GTTAGAGPRC
     201 RPINATLAVE KEGCPVCITV NTTICAGYCP TMTRVLQGVL PALPQVVCNY
         251 RDVRFESIRL PGCPRGVNPV VSYAVALSCQ CALCRRSTTD CGGPKDHPLT
         301 CDDPRFQDSS SSKAPPPSLP SPSRLPGPSD TPILPQ
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191-331 HITS AT: MFUnspecified CI MAN SR CA LCCA, CAPLUS, TOXCENTER, USPATFULL STN Files: 2 REFERENCES IN FILE CA (1962 TO DATE) 2 REFERENCES IN FILE CAPLUS (1962 TO DATE) REFERENCE 1: 134:188974 REFERENCE 2: 127:244008 L28 ANSWER 6 OF 7 REGISTRY COPYRIGHT 2002 ACS RN **195460-70-9** REGISTRY CN 20-161-Tumor necrosis factor receptor p55 (human clone pSVL-hTBP1.hCG.beta.) fusion protein with peptide (synthetic linker) fusion protein with chorionic gonadotropin (human .beta.-subunit fragment) (9CI) (CA INDEX NAME) OTHER NAMES: 4: PN: US6194177 SEQID: 4 claimed protein CN FS PROTEIN SEQUENCE SOL 307 PATENT ANNOTATIONS (PNTE): Sequence | Patent Source | Reference ======+======== Not Given | US6194177 |claimed |SEQID 4 SEO 1 SRTSLLLAFG LLCLPWLQEG SADSVCPQGK YIHPQNNSIC CTKCHKGTYL 51 YNDCPGPGQD TDCRECESGS FTASENHLRH CLSCSKCRKE MGQVEISSCT 101 VDRDTVCGCR KNQYRHYWSE NLFQCFNCSL CLNGTVHLSC QEKQNTVCTC 151 HAGFFLRENE CVSCAGAGPR CRPINATLAV EKEGCPVCIT VNTTICAGYC 201 PTMTRVLQGV LPALPQVVCN YRDVRFESIR LPGCPRGVNP VVSYAVALSC 251 OCALCRRSTT DCGGPKDHPL TCDDPRFODS SSSKAPPPSL PSPSRLPGPS 301 DTPILPO HITS AT: 162-302 MF Unspecified CI MAN SR CA LC CA, CAPLUS, TOXCENTER, USPATFULL STN Files: 2 REFERENCES IN FILE CA (1962 TO DATE) 2 REFERENCES IN FILE CAPLUS (1962 TO DATE) REFERENCE 1: 134:188974 REFERENCE 2: 127:244008 ANSWER 7 OF 7 REGISTRY COPYRIGHT 2002 ACS RN **76050-53-8** REGISTRY CN Gonadotropin, chorionic pre- (human .beta.-subunit protein moiety reduced) (9CI) (CA INDEX NAME) OTHER NAMES: CN 10: PN: WO0041717 FIGURE: 1A unclaimed protein CN Gonadotropin, chorionic (human embryo .beta.-subunit)

FS PROTEIN SEQUENCE SOL 165

PATENT ANNOTATIONS (PNTE):

Sequence | Patent Source | Reference

Not Given|WO2000041717

|unclaimed |FIGURE 1A

SEQ 1 MEMFQGLLLL LLLSMGGTWA SKEPLRPRCR PINATLAVEK EGCPVCITVN

51 TTICAGYCPT MTRVLQGVLP ALPQVVCNYR DVRFESIRLP GCPRGVNPVV

51 IIICAGICPI MIRVLQGVLP ALPQVVCNIR DVRFESIRLP GCPRGVNPVV

101 SYAVALSCQC ALCRRSTTDC GGPKDHPLTC DDPRFQDSSS SKAPPPSLPS

151 PSRLPGPSDT PILPO

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HITS AT: 20-160

RELATED SEQUENCES AVAILABLE WITH SEQLINK

MF Unspecified

CI MAN

LC STN Files: CA, CAPLUS, TOXCENTER, USPATFULL

6 REFERENCES IN FILE CA (1962 TO DATE)

6 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 133:134164

REFERENCE 2: 132:246924

REFERENCE 3: 116:208630

REFERENCE 4: 100:97541

REFERENCE 5: 99:207297

REFERENCE 6: 94:11741

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FILE 'HCAPLUS' ENTERED AT 17:11:02 ON 11 OCT 2002
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FILE COVERS 1907 - 11 Oct 2002 VOL 137 ISS 16 FILE LAST UPDATED: 10 Oct 2002 (20021010/ED)

This file contains CAS Registry Numbers for easy and accurate

substance identification.

CAS roles have been modified effective December 16, 2001. Please check your SDI profiles to see if they need to be revised. For information on CAS roles, enter HELP ROLES at an arrow prompt or use the CAS Roles thesaurus (/RL field) in this file.

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    ANSWER 1 OF 11 HCAPLUS COPYRIGHT 2002 ACS
ΑN
     2001:843811 HCAPLUS
     136:1103
DN
     Treatment and prevention of HIV infection by administration of derivatives
TΤ
     of human chorionic gonadotropin
IN
     Gallo, Robert C.; Bryant, Joseph; Lunardi-Iskandar, Yanto
PA
     University of Maryland Biotechnology Institute, USA
SO
     U.S., 51 pp., Cont.-in-part of U.S. Ser. No. 669,681, abandoned.
    CODEN: USXXAM
DT
    Patent
LA
    English
    ICM A61K039-00
IC
NCL
    424198100
CC
     2-4 (Mammalian Hormones)
     Section cross-reference(s): 63
FAN.CNT 2
                    KIND DATE
    PATENT NO.
                                         APPLICATION NO. DATE
                     ----
     ______
                                          ______
PΙ
    US 6319504
                    B1
                           20011120
                                          US 1996-709948 19960909 <--
    WO 9749373 A2 19971231
WO 9749373 A3 19980226
                                          WO 1997-US11202 19970624 <--
            AL, AM, AU, AZ, BA, BB, BG, BR, BY, CA, CN, CU, CZ, EE, GE, GH,
            HU, IL, IS, JP, KG, KP, KR, KZ, LC, LK, LR, LT, LV, MD, MG, MK,
            MN, MX, NO, NZ, PL, RO, RU, SG, SI, SK, SL, TJ, TM, TR, TT, UA,
             US, US, UZ, VN, YU, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
        RW: GH, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, DE, DK, ES, FI, FR,
            GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA,
             GN, ML, MR, NE, SN, TD, TG
    AU 9738792
                     A1
                          19980114
                                          AU 1997-38792
                                                          19970624 <--
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IE, FI PRAI US 1996-669681 B2 19960624 <--US 1996-709948 A2 19960909 <--WO 1997-US11202 W 19970624 <--

A2

19990908

EP 939589

The present invention relates to .beta.-hCG, particularly .beta.-hCG AB proteins having a sequence of amino acids 41-54, 45-54, 47-53, 45-57 and 45-58 and analogs and derivs. thereof. The invention further relates to methods of treatment and prevention of HIV infection by administration of a therapeutic compd. of the invention. The peptides of the invention can also be used to treat Kaposi's sarcoma and hemopoiesis dysfunction. Such therapeutic compds. include hCG, .beta.-hCG and .beta.-hCG peptides, analogs and derivs. of hCG, .beta.-hCG and .beta.-hCG peptides, and nucleic acids encoding hCG, .beta.-hCG and .beta.-hCG peptides. In a preferred embodiment, .beta.-hCG peptides, particularly .beta.-hCG peptides of amino acids 47-53, 45-57 or 45-58 are administered to a subject for treatment or prevention of HIV infection in that subject. invention also provides methods for screening hCG prepns. for activity in treating or preventing HIV infection. Pharmaceutical compns. and methods of administration of therapeutics are also provided.

EP 1997-936023

AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,

19970624 <--

- HIV infection treatment chorionic gonadotropin deriv; Kaposis sarcoma STtreatment chorionic gonadotropin deriv; hemopoiesis dysfunction treatment chorionic gonadotropin deriv
- ΙT Sarcoma

(Kaposi's, inhibitors; treatment and prevention of HIV infection, Kaposi's sarcoma, and hemopoiesis dysfunction by administration of derivs. of human chorionic gonadotropin)

IT Hematopoiesis

(prohematopoietic effects; treatment and prevention of HIV infection, Kaposi's sarcoma, and hemopoiesis dysfunction by administration of derivs. of human chorionic gonadotropin)

IT Drug delivery systems

(treatment and prevention of HIV infection by administration of derivs. of human chorionic gonadotropin)

IT Chemokines

Macrophage inflammatory protein 1.alpha.

Macrophage inflammatory protein 1.beta.

RANTES (chemokine)

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(treatment and prevention of HIV infection by administration of derivs. of human chorionic gonadotropin in combination with a chemokine)

IT Gene therapy

(treatment and prevention of HIV infection by administration of nucleic acids encoding .beta.-hCG or .beta.-hCG peptides)

IT Anti-AIDS agents

(treatment and prevention of HIV infection, Kaposi's sarcoma, and hemopoiesis dysfunction by administration of derivs. of human chorionic gonadotropin)

IT 201351-22-6

RL: PRP (Properties)

(Unclaimed; treatment and prevention of HIV infection by administration of derivs. of human chorionic gonadotropin)

IT 7481-89-2, DdC 30516-87-1, AZT 69655-05-6, Didanosine 127779-20-8, Saquinavir 134678-17-4, 3TC

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(treatment and prevention of HIV infection by administration of derivs. of human chorionic gonadotropin in combination with another antiviral agent)

ŦΤ 108303-18-0 163007-06-5 201350-97-2 201351-01-1 201351-02-2 201351-03-3 201351-04-4 201351-05-5 201351-06-6 201351-07-7 201351-09-9 201351-13-5 201351-18-0 201351-19-1 201351-20-4 201351-21-5 201351-23-7 201351-24-8 201351-55-5 201492-48-0 201492-49-1 374728-54-8

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(treatment and prevention of HIV infection, Kaposi's sarcoma, and hemopoiesis dysfunction by administration of derivs. of human chorionic gonadotropin)

IT 202017-03-6

RL: PRP (Properties)

(unclaimed nucleotide sequence; treatment and prevention of HIV infection by administration of derivs. of human chorionic gonadotropin)

IT **202016-40-8** 375375-79-4

RL: PRP (Properties)

(unclaimed protein sequence; treatment and prevention of HIV infection by administration of derivs. of human chorionic gonadotropin)

RE.CNT 132 THERE ARE 132 CITED REFERENCES AVAILABLE FOR THIS RECORD RE

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AN
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DN
    135:1276
TI
    Chimeric genes for single-chain forms of glycoprotein hormones and their
     expression in host cells
IN
    Biome, Irving; Moyle, William R.
PΑ
    Washington University, USA
    U.S., 87 pp., Cont.-in-part of U.S. 853,524.
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     3-3 (Biochemical Genetics)
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    The DNA encoding single-chain forms of the glycoprotein hormones LH, FSH,
AB
    TSH, and CG are disclosed. The .alpha. and .beta. subunits of the
    wild-type heterodimers or their variants or their fragments are covalently
    linked, optionally through a linker moiety. Some of the single-chain
     forms are agonists and others antagonists of the glycoprotein hormone
    activity. The DNA for these fusion proteins are expressed in host cells
     in order to produce the hormone derivs.
ST
    sequence human single chain LH FSH TSH CG gene
ΙT
    Molecular cloning
        (chimeric genes for single-chain forms of glycoprotein hormones and
       their expression in host cells)
IT
    Chimeric gene
     RL: BUU (Biological use, unclassified); PRP (Properties); BIOL (Biological
     study); USES (Uses)
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(chimeric genes for single-chain forms of glycoprotein hormones and
        their expression in host cells)
ΙT
     DNA sequences
         (of chimeric genes for single-chain forms of human glycoprotein
        hormones)
IT
     Protein sequences
         (of single-chain forms of human glycoprotein hormones)
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ΤТ
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                                            9002-67-9, LH
                                                             9002-68-0, FSH
     9002-71-5, TSH
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
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                                                     342058-66-6P
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RE.CNT
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               THERE ARE 20 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE
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(2) Anon; Chemical Abstracts 1982, V97(17), P94
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- L26 ANSWER 3 OF 11 HCAPLUS COPYRIGHT 2002 ACS
- AN 2001:145198 HCAPLUS
- DN 134:188974
- TI DNA encoding human hybrid heterodimeric proteins for modulation of protein-protein interactions
- IN Campbell, Robert K.; Jameson, Bradford A.; Chappel, Scott C.
- PA Applied Research Systems ARS Holding N.V., Neth. Antilles
- SO U.S., 35 pp., Cont.-in-part of U.S. Ser. No. 804,166. CODEN: USXXAM
- DT Patent
- LA English
- IC C12P021-04
- NCL 435069700
- CC 3-2 (Biochemical Genetics)

Section cross-reference(s): 1, 2, 13

FAN.CNT 2

IT

Animal cell line

1174.0141 2						
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	US 6193972	В1	20010227		US 1997-804166	19970220 <
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- This invention relates to a hybrid protein of two amino acid sequences AB joined directly or with a peptide linker. Each hybrid protein sequence contains the binding portion of a receptor, such as tumor necrosis factor receptor 1 (TBP1), or a ligand linked to a subunit of a heterodimeric proteinaceous hormone, such as human chorionic gonadotropin (hCG). Each hybrid protein sequence contains a corresponding hormone subunit so as to form a heterodimer upon coexpression. Corresponding DNA mols., expression vectors, host cells, and a method of producing such proteins are claimed. These hybrid proteins could result in monofunctional, bifunctional, or multifunctional mols. for modulating protein-protein interactions, for example by sequestering ligands or regulating receptor activity. Recombinant fusion proteins TBP1-hCG(.alpha./.beta.) were produced, secreted into culture media of transfected mammalian cells, and formed heterodimers. The TBP1-hCG(.alpha./.beta.) proteins inhibited tumor necrosis factor cytotoxicity in a bioassay using the human breast carcinoma cell line BT-20. A plasmid was constructed for expression of the FSH .beta. subunit fused to the extracellular domain of the FSH receptor with a thrombin cleavage site and thrombin receptor extracellular tethering domain.
- ST recombinant DNA expression fusion protein heterodimeric receptor ligand hormone; tumor necrosis factor receptor chorionic gonadotropin fusion protein bioassay; plasmid FSH FSHR fusion cleavable peptide linker
 - (BT20; modulation of protein-protein interactions by human hybrid heterodimeric tumor necrosis factor receptor 1-human chorionic

gonadotropin proteins measured by bioassay) TT Animal cell line (CHO; recombinant expression of human hybrid heterodimeric proteins, for modulation of protein-protein interactions) Plasmid vectors TΤ (CMV/FSHR-EC/TR/FSH.beta.; DNA encoding human hybrid heterodimeric proteins for modulation of protein-protein interactions) TΤ Animal cell line (COS-7; recombinant expression of human hybrid heterodimeric proteins, for modulation of protein-protein interactions) ΤТ Protein receptors RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (DNA encoding human hybrid heterodimeric proteins contq. a protein receptor, for modulation of protein-protein interactions,) ΙT Molecular association Molecular cloning (DNA encoding human hybrid heterodimeric proteins for modulation of protein-protein interactions) IT Chimeric gene Fusion proteins (chimeric proteins) RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (DNA encoding human hybrid heterodimeric proteins for modulation of protein-protein interactions) Primers (nucleic acid) TT RL: ARG (Analytical reagent use); BUU (Biological use, unclassified); PRP (Properties); ANST (Analytical study); BIOL (Biological study); USES (Uses) (DNA; PCR primers used for construction of DNA encoding human hybrid heterodimeric proteins for modulation of protein-protein interactions) cDNA sequences IT (encoding human hybrid heterodimeric proteins for modulation of protein-protein interactions) IT FSH receptors RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (extracellular domain; DNA encoding human hybrid heterodimeric proteins contg. FSH receptor, for modulation of protein-protein interactions) IT Thrombin receptors RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (extracellular domain; DNA encoding human hybrid heterodimeric proteins contg. thrombin receptor, for modulation of protein-protein interactions) Proteins, specific or class ΙT RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (ligand-binding; DNA encoding human hybrid heterodimeric proteins contq. a ligand-binding protein, for modulation of protein-protein interactions) ΙT Proteins, specific or class RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (ligands; DNA encoding human hybrid heterodimeric proteins for modulation of protein-protein interactions) TTPeptides, biological studies RL: BUU (Biological use, unclassified); PRP (Properties); BIOL (Biological study); USES (Uses) (linker; DNA encoding human hybrid heterodimeric proteins contg. a linker peptide, for modulation of protein-protein interactions) IT Cytoprotective agents Cytotoxicity (modulation of protein-protein interactions by human hybrid heterodimeric tumor necrosis factor receptor 1-human chorionic gonadotropin proteins measured by bioassay) ΙT Protein sequences (of human hybrid heterodimeric proteins for modulation of

protein-protein interactions)

Tumor necrosis factor receptors IΤ RL: BAC (Biological activity or effector, except adverse); BPN (Biosynthetic preparation); BPR (Biological process); BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study); PREP (Preparation); PROC (Process) (p55, fusion products; DNA encoding human hybrid heterodimeric proteins contg. tumor necrosis factor receptor p55, for modulation of protein-protein interactions) IT Plasmid vectors (pSVL-based and D.alpha.-based; DNA encoding human hybrid heterodimeric proteins for modulation of protein-protein interactions) IT DNA RL: ARG (Analytical reagent use); BUU (Biological use, unclassified); PRP (Properties); ANST (Analytical study); BIOL (Biological study); USES (primer; PCR primers used for construction of DNA encoding human hybrid heterodimeric proteins for modulation of protein-protein interactions) IT Enzymes, biological studies RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses) (protein-degrading, proteolytic cleavage site; DNA encoding human hybrid heterodimeric proteins contg. a proteolytic cleavage site, for modulation of protein-protein interactions) ΙT Hormones, animal, biological studies RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (protein; DNA encoding human hybrid heterodimeric proteins, including hormones, for modulation of protein-protein interactions,) ΙT Secretion (process) (protein; modulation of protein-protein interactions by secreted human hybrid heterodimeric tumor necrosis factor receptor 1-human chorionic gonadotropin proteins measured by bioassay) ΙT Animal cell line (recombinant expression of human hybrid heterodimeric proteins, for modulation of protein-protein interactions) IT 69287-89-4 RL: PRP (Properties) (Unclaimed; dNA encoding human hybrid heterodimeric proteins for modulation of protein-protein interactions) ΤТ 195460-68-5P **195460-70-9P** 195460-72-1P **195460-74-3P** RL: BAC (Biological activity or effector, except adverse); BPN (Biosynthetic preparation); BPR (Biological process); BSU (Biological study, unclassified); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); PROC (Process); USES (Uses) (amino acid sequence; of human hybrid heterodimeric proteins for modulation of protein-protein interactions) IΤ 9002-04-4, Thrombin RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses) (cleavage site; DNA encoding human hybrid heterodimeric proteins contg. a thrombin cleavage site, for modulation of protein-protein interactions) ΙT 195460-69-6 195460-73-2 328049-26-9 328049-27-0 RL: BSU (Biological study, unclassified); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (nucleotide sequence; DNA encoding human hybrid heterodimeric proteins for modulation of protein-protein interactions) 328053-44-7 IT 328053-41-4 328053-42-5 328053-43-6 328053-45-8 328053-48-1 328053-49-2 328053-50-5 328053-46-9 328053-47-0 328053-52-7 328053-51-6 RL: PRP (Properties) (unclaimed nucleotide sequence; dNA encoding human hybrid heterodimeric proteins for modulation of protein-protein interactions) IT9002-61-3DP, Human chorionic gonadotropin, fusion products

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RL: BPN (Biosynthetic preparation); BPR (Biological process); BSU
     (Biological study, unclassified); PRP (Properties); THU (Therapeutic use);
     BIOL (Biological study); PREP (Preparation); PROC (Process); USES (Uses)
        (.alpha. and .beta. subunits; DNA encoding human hybrid heterodimeric
        proteins contg. chorionic gonadotropin subunits, for modulation of
        protein-protein interactions)
     9002-68-0D, Follicle stimulating hormone, fusion products
ΙT
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (.beta. subunit; DNA encoding human hybrid heterodimeric proteins
        contg. FSH, for modulation of protein-protein interactions)
RE.CNT
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     ANSWER 4 OF 11 HCAPLUS COPYRIGHT 2002 ACS
     1999:670049 HCAPLUS
AN
DN
     131:295923
TI
     Method of promoting hematopoiesis using cyclic peptides derived from human
     chorionic gonadotropin fragments
ΙN
     Gallo, Robert C.; Bryant, Joseph; Lunardi-Iskandar, Yanto
PA
     University of Maryland Biotechnology Institute, USA
SO
     U.S., 40 pp., Cont.-in-part of U.S. Ser. No. 669,654, anbandoned.
     CODEN: USXXAM
DT
     Patent
LA
     English
IC
     ICM A61K038-12
     ICS C07K007-64
NCL
     424185100
     2-4 (Mammalian Hormones)
CC
     Section cross-reference(s): 63
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                      W
                            19970624
                                     <--
AB
     The present invention relates to methods of treating or preventing
     diseases or disorders assocd. with hematopoietic deficiency by
     administration of cyclic peptides derived from human .beta.-human
     chorionic gonadotropin fragments. The invention also relates to methods
     of treating or preventing diseases or disorders assocd. with hematopoietic
     deficiency by administration of hematopoietic cells, the nos. of which
     have been increased by contacting the cells with human chorionic
     gonadotropin, .beta.-human chorionic gonadotropin or a peptide contg. a
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sequence of a portion of .beta.-human chorionic gonadotropin. The

invention also provides assays for the utility of particular human chorionic gonadotropin prepns. in the treatment or prevention of hematopoietic deficiencies or in the increasing of hematopoietic cell nos. in vitro. Pharmaceutical compns. and methods of administration of are also provided.

ST hematopoiesis promotion human chorionic gonadotropin deriv

IT Hematopoietic precursor cell

(method of promoting hematopoiesis by administration of hematopoietic cells, the nos. of which have been increased by contacting the cells with cyclic peptides derived from .beta.-human chorionic gonadotropin fragments)

IT Drug delivery systems

Hematopoiesis

(method of promoting hematopoiesis using cyclic peptides derived from human chorionic gonadotropin fragments)

IT 202016-40-8DP, fragments, cyclic peptides derived from
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); PUR (Purification or recovery); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(method of promoting hematopoiesis using cyclic peptides derived from human chorionic gonadotropin fragments)

IT 72979-70-5D, cyclic peptides derived from 163007-06-5D, cyclic peptides 201350-97-2D, cyclic peptides derived from derived from 201351-01-1D, 201351-02-2D, cyclic peptides derived from cyclic peptides derived from 201351-03-3D, cyclic peptides derived from 201351-04-4D, cyclic peptides 201351-05-5D, cyclic peptides derived from derived from 201351-06-6D, cyclic peptides derived from \cdot 201351-07-7D, cyclic peptides derived from 201351-09-9D, cyclic peptides derived from 201351-13-5D, cyclic peptides 201351-18-0D, cyclic peptides derived from derived from 201351-19-1D, cyclic peptides derived from 201351-20-4D, cyclic peptides derived from 201351-21-5D, cyclic peptides derived from 201351-22-6D, cyclic peptides derived from 201351-23-7D, cyclic peptides derived from 201351-24-8D, cyclic peptides derived from 201351-55-5 201492-48-0D, cyclic peptides derived from 201492-49-1D, cyclic peptides derived from RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES

(method of promoting hematopoiesis using cyclic peptides derived from human chorionic gonadotropin fragments)

IT 202017-03-6

RE.CNT

RE

RL: PRP (Properties)

(unclaimed nucleotide sequence; method of promoting hematopoiesis using
cyclic peptides derived from human chorionic gonadotropin fragments)
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L26 ANSWER 5 OF 11 HCAPLUS COPYRIGHT 2002 ACS
AN
     1998:42294 HCAPLUS
DN
     128:124125
ΤI
    Methods of promoting hematopoiesis using derivatives of human chorionic
     gonadotropin
ΙN
     Gallo, Robert C.; Bryant, Joseph; Lunardi-Iskandar, Yanto
PA
     University of Maryland Biotechnology Institute, USA; Gallo, Robert C.;
     Bryant, Joseph; Lunardi-Iskandar, Yanto PCT Int. Appl., 175 pp.
SO
     CODEN: PIXXD2
DT
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     English
IC
     ICM A61K038-00
     ICS
         C07K001-00; C12N015-00
     2-4 (Mammalian Hormones)
     Section cross-reference(s): 63
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             MN, MX, NO, NZ, PL, RO, RU, SG, SI, SK, SL, TJ, TM, TR, TT, UA,
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     US 1996-709924
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     WO 1997-US11209
                            19970624
     The present invention relates to methods of treating or preventing
AB
     diseases or disorders assocd. with hematopoietic deficiency by
     administration of human chorionic gonadotropin, .beta.-human chorionic
     gonadotropin, a peptide contg. a sequence of one or more portions of
     .beta.-human chorionic gonadotropin, or fractions of a source of native
     human chorionic gonadotropin or native .beta.-human chorionic
     gonadotropin. The invention also relates to methods of treating and
     preventing diseases or disorders assocd. with hematopoietic deficiency by
     administration of hematopoietic cells, the nos. of which have been
     increased by contacting the cells with a therapeutic of the invention.
     Pharmaceutical compns. and methods of administration are also provided.
ST
     hematopoiesis promotion human chorionic gonadotropin deriv
TΤ
     Bone marrow
        (cells; treating and preventing diseases or disorders assocd. with
        hematopoietic deficiency by administration of hematopoietic cells whose
        nos. have been increased by contacting them with derivs. of human
        chorionic gonadotropin)
IT
     Fusion proteins (chimeric proteins)
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
     (Uses)
        (contg. .beta.-hCG fragments; methods of promoting hematopoiesis using
        derivs. of human chorionic gonadotropin)
TΤ
     Peptides, biological studies
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
     (Uses)
        (cyclic; methods of promoting hematopoiesis using derivs. of human
        chorionic gonadotropin)
     CD4-positive T cell
IT
     Simian immunodeficiency virus
        (drug screening for human chorionic gonadotropin derivs. with
        prohematopoietic activity using CD4+ T cells in an SIV infected monkey)
TΤ
     Urine
        (early pregnancy urine as a source of hCG fractions; methods of
        promoting hematopoiesis using derivs. of human chorionic gonadotropin)
TT
     Drug screening
        (for pro-hematopoietic activity; methods of promoting hematopoiesis
        using derivs. of human chorionic gonadotropin)
ΙT
     Liquid chromatography
        (gel filtration sizing column chromatog. for fractionation of native
        hCG and native .beta.-hCG; methods of promoting hematopoiesis using
        derivs. of human chorionic gonadotropin)
TT
     Purpura (disease)
        (idiopathic thrombocytopenic; methods of promoting hematopoiesis using
        derivs. of human chorionic gonadotropin in patients with HIV infection,
        idiopathic thrombocytopenic purpura, anemia, or neutropenia)
```

(methods of promoting hematopoiesis using derivs. of human chorionic

IT Anemia (disease)

Hematopoiesis

gonadotropin)

ΙT

Human immunodeficiency virus 1

(methods of promoting hematopoiesis using derivs. of human chorionic gonadotropin in patients with HIV infection, idiopathic thrombocytopenic purpura, anemia, or neutropenia)

IT Antitumor agents

Radiotherapy

(methods of promoting hematopoiesis using derivs. of human chorionic gonadotropin in subjects undergoing chemotherapy or radiation therapy)

IT Drug delivery systems

(methods of promoting hematopoiesis using pharmaceutical formulations contg. derivs. of human chorionic gonadotropin)

IT Agranulocytosis

(neutropenia; methods of promoting hematopoiesis using derivs. of human chorionic gonadotropin in patients with HIV infection, idiopathic thrombocytopenic purpura, anemia, or neutropenia)

IT Fractionation

(of native hCG and native .beta.-hCG; methods of promoting hematopoiesis using derivs. of human chorionic gonadotropin)

IT Embryo, animal

(stem cell; treating and preventing diseases or disorders assocd. with hematopoietic deficiency by administration of hematopoietic cells whose nos. have been increased by contacting them with derivs. of human chorionic gonadotropin)

IT Hematopoietic precursor cell

(stem; treating and preventing diseases or disorders assocd. with hematopoietic deficiency by administration of hematopoietic cells whose nos. have been increased by contacting them with derivs. of human chorionic gonadotropin)

IT Blood cell

(treating and preventing diseases or disorders assocd. with hematopoietic deficiency by administration of hematopoietic cells whose nos. have been increased by contacting them with derivs. of human chorionic gonadotropin)

IT Gene therapy

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(using nucleic acids encoding human chorionic gonadotropin derivs.; methods of promoting hematopoiesis using derivs. of human chorionic gonadotropin)

IT Infection

201352-37-6

201423-37-2

(viral; methods of promoting hematopoiesis using derivs. of human chorionic gonadotropin in patients with HIV infection, idiopathic thrombocytopenic purpura, anemia, or neutropenia)

IT 9002-61-3P, Human chorionic gonadotropin 202016-40-8P
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); PUR (Purification or recovery); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(methods of promoting hematopoiesis using derivs. of human chorionic gonadotropin)

gonadotropin) IT 9002-61-3D, Human chorionic gonadotropin, fragments, analogs, and derivs. 72979-70-5 108303-18-0 108303-18-0D, analogs 163007-06-5 201351-02-2 201350-97-2 201351-01-1 201351-03-3 201351-04-4 201351-09-9 201351-05-5 201351-06-6 201351-07-7 201351-13-5 201351-18-0 201351-19-1 201351-20-4 201351-21-5 201351-22-6 201351-29-3 201351-23-7 201351-24-8 201351-26-0 201351-28-2 201351-30-6 201351-31-7 201351-33-9 201351-34-0 201351-35-1 201351-37-3 201351-38-4 201351-40-8 201351-41-9 201351-55-5 201351-70-4 201351-56-6 201351-60-2 201351-74-8 201351-77-1 201351-82-8 201351-86-2 201351-89-5 201351-92-0 201351-95-3 201351-98-6 201352-01-4 201352-05-8 201352-13-8 201352-16-1 201352-24-1 201352-27-4 201352-30-9 201352-33-2 201352-36-5

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201492-48-0 201492-49-1 201492-50-4 201492-51-5 202016-40-8D , fragments, analogs, and derivs. 202017-03-6 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (methods of promoting hematopoiesis using derivs. of human chorionic gonadotropin) ANSWER 6 OF 11 HCAPLUS COPYRIGHT 2002 ACS 1998:42256 HCAPLUS 128:124124 Treatment and prevention of HIV infection by administration of derivatives of human chorionic gonadotropin Gallo, Robert C.; Bryant, Joseph; Lunardi-Iskandar, Yanto University of Maryland Biotechnology Institute, USA; Gallo, Robert C.; Bryant, Joseph; Lunardi-Iskandar, Yanto PCT Int. Appl., 173 pp. CODEN: PIXXD2 Patent English ICM A61K 2-4 (Mammalian Hormones) Section cross-reference(s): 63 FAN.CNT 2 PATENT NO. KIND DATE APPLICATION NO. DATE ----_____ WO 9749373 A2 19971231 WO 1997-US11202 19970624 <--WO 9749373 A3 19980226 AL, AM, AU, AZ, BA, BB, BG, BR, BY, CA, CN, CU, CZ, EE, GE, GH, HU, IL, IS, JP, KG, KP, KR, KZ, LC, LK, LR, LT, LV, MD, MG, MK, MN, MX, NO, NZ, PL, RO, RU, SG, SI, SK, SL, TJ, TM, TR, TT, UA, US, US, UZ, VN, YU, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG US 6319504 В1 20011120 US 1996-709948 19960909 <--AU 9738792 A1 19980114 AU 1997-38792 19970624 <--EP 939589 A2 19990908 EP 1997-936023 19970624 <--AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, PRAI US 1996-669681 A2 19960624 <--US 1996-709948 A2 19960909 <--WO 1997-US11202 W 19970624 <--The present invention relates to .beta.-hCG, particularly certain .beta.-hCG peptides, and analogs and derivs. thereof. The invention also relates to fractions of a source of native hCG or native .beta.-hCG, which fractions are active in inhibiting HIV infection or replication, against Kaposi's sarcoma or have a pro-hematopoietic effect. The invention further relates to methods of treatment and prevention of HIV infection by administration of a therapeutic compd. of the invention. Such therapeutic compds. include hCG, .beta.-hCG and .beta.-hCG peptides, analogs and derivs. of hCG, .beta.-hCG and .beta.-hCG peptides, and nucleic acids encoding hCG, .beta.-hCG and .beta.-hCG peptides, and therapeutically and prophylactically effective fractions of sources of native hCG or native .beta.-hCG. Pharmaceutical compns. and methods of administration of therapeutics are also provided. HIV infection treatment beta chorionic gonadotropin Sarcoma (Kaposi's; treatment of Kaposi's sarcoma by administration of derivs. of human chorionic gonadotropin) Antigens

RL: BPR (Biological process); BSU (Biological study, unclassified); BUU (Biological use, unclassified); BIOL (Biological study); PROC (Process); USES (Uses)

(SIV p27; drug screening of .beta.-hCG or its fractions for anti-HIV activity using HIV-1 p24 antigen, HIV-1 LTR, HIV-1 derived RNA transcripts, or SIV p27 antigen)

IT Peptides, biological studies

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(cyclic; treatment and prevention of HIV infection by administration of derivs. of human chorionic gonadotropin)

IT Drug screening

Simian immunodeficiency virus

(drug screening of .beta.-hCG or its fractions for anti-HIV activity using HIV-1 p24 antigen, HIV-1 LTR, HIV-1 derived RNA transcripts, or SIV p27 antigen)

IT mRNA

RL: BPR (Biological process); BSU (Biological study, unclassified); BUU (Biological use, unclassified); BIOL (Biological study); PROC (Process); USES (Uses)

(drug screening of .beta.-hCG or its fractions for anti-HIV activity using HIV-1 p24 antigen, HIV-1 LTR, HIV-1 derived RNA transcripts, or SIV p27 antigen)

IT Urine

(early pregnancy urine as a source of hCG fractions; treatment and prevention of HIV infection by administration of derivs. of human chorionic gonadotropin)

IT Chemokines

Macrophage inflammatory protein 1.alpha.

Macrophage inflammatory protein 1.beta.

RANTES (chemokine)

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(fusion proteins, with .beta.-hCG fragments; treatment and prevention of HIV infection by administration of derivs. of human chorionic gonadotropin)

IT Liquid chromatography

(gel filtration sizing column chromatog. for fractionation of native hCG and native .beta.-hCG; treatment and prevention of HIV infection by administration of derivs. of human chorionic gonadotropin or nucleic acids encoding the derivs.)

IT Genetic element

RL: BPR (Biological process); BSU (Biological study, unclassified); BUU (Biological use, unclassified); BIOL (Biological study); PROC (Process); USES (Uses)

(long terminal repeat; drug screening of .beta.-hCG or its fractions for anti-HIV activity using HIV-1 p24 antigen, HIV-1 LTR, HIV-1 derived RNA transcripts, or SIV p27 antigen)

IT Fractionation

(of native hCG and native .beta.-hCG; treatment and prevention of HIV infection by administration of derivs. of human chorionic gonadotropin)

IT Antigens

RL: BPR (Biological process); BSU (Biological study, unclassified); BUU (Biological use, unclassified); BIOL (Biological study); PROC (Process); USES (Uses)

(p24; drug screening of .beta.-hCG or its fractions for anti-HIV activity using HIV-1 p24 antigen, HIV-1 LTR, HIV-1 derived RNA transcripts, or SIV p27 antigen)

IT Hematopoiesis

(pro-hematopoietic activity of derivs. of human chorionic gonadotropin)

IT Antitumor agents

(sarcoma; treatment of Kaposi's sarcoma by administration of derivs. of human chorionic gonadotropin)

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ΙT
     Antiviral agents
     Human immunodeficiency virus 1
        (treatment and prevention of HIV infection by administration of derivs.
        of human chorionic gonadotropin)
TΤ
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
     (Uses)
        (treatment and prevention of HIV infection by administration of derivs.
        of human chorionic gonadotropin or nucleic acids encoding the derivs.)
ΙT
     Drug delivery systems
        (treatment and prevention of HIV infection by administration of
        formulations contg. derivs. of human chorionic gonadotropin)
TΤ
     Fusion proteins (chimeric proteins)
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
        (.beta.-hCG fragment joined to a protein different from .beta.-hCG;
        treatment and prevention of HIV infection by administration of derivs.
        of human chorionic gonadotropin)
TΤ
     9002-61-3P, Human chorionic gonadotropin 202016-40-8P
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); PUR (Purification or recovery); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (treatment and prevention of HIV infection by administration of derivs.
        of human chorionic gonadotropin)
TT
     9002-61-3D, Human chorionic gonadotropin, fragments, analogs, and derivs.
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     48-145-Gonadotropin, chorionic (human .beta.-subunit)
                                                              201492-49-1,
     58-145-Gonadotropin, chorionic (human .beta.-subunit)
                                                              201492-51-5
     201688-15-5
                   201688-16-6 202016-40-8D, fragments, analogs, and
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               202017-03-6
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     study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
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        (treatment and prevention of HIV infection by administration of derivs.
        of human chorionic gonadotropin)
L26 ANSWER 7 OF 11 HCAPLUS COPYRIGHT 2002 ACS
     1997:568294 HCAPLUS
ΑN
DN
     127:244008
ΤI
     Recombinant fusion proteins comprising ligand-binding receptor fragment
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IN Campbell, Robert K.; Jameson, Bradford A.; Chappel, Scott C.

linked with hormone subunit, heterodimer formation, and pharmaceutical

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Applied Research Systems ARS Holding N.V., Neth. Antilles; Campbell,
PA
    Robert K.; Jameson, Bradford A.; Chappel, Scott C.
    PCT Int. Appl., 60 pp.
SO
    CODEN: PIXXD2
DT
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LA
    English
    ICM C12N015-62
TC
    ICS C12N015-16; C07K014-59; C07K014-715; C07K014-72; C07K016-46;
         C12N015-85; C12N005-10; A61K038-24
CC
     3-2 (Biochemical Genetics)
     Section cross-reference(s): 1, 2, 13, 15
FAN.CNT 2
                     KIND DATE
    PATENT NO.
                                          APPLICATION NO. DATE
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    WO 9730161
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                     W
                           19970220 <--
    WO 1997-US2315
AΒ
    This invention comprises a hybrid protein including two coexpressed amino
    acid sequences which form a heterodimer. Each sequence contains the
    binding portion of a receptor, such as tumor necrosis factor binding
    protein TBP1 or TBP2, or a ligand, such as interleukin-6,
    interferon-.beta., or thrombopoietin (TPO), linked to a subunit of a
    heterodimeric proteinaceous hormone, such as human chorionic gonadotropin.
    Each coexpressed sequence contains a corresponding hormone subunit so as
    to form a heterodimer upon expression. Corresponding DNA mols.,
    expression vectors and host cells are also disclosed as are pharmaceutical
    compns. and a method of producing such proteins. The general method is
    exemplified by TBP1(20-161) fusion products with human chorionic
    gonadotropin .alpha. subunit coexpression with TBP1(20-161) fusion
    products with human chorionic gonadotropin .beta. subunit. The hybrid
    proteins were coexpressed by COS-7 cells, formed heterodimers, and
    protected BT-20 cells against TNF.alpha.-induced cytotoxicity.
    receptor fusion hormone subunit recombinant heterodimer; ligand fusion
ST
    hormone subunit recombinant heterodimer; chorionic gonadotropin subunit
    fusion TBP protein; TNF binding protein fusion hormone subunit; tumor
    necrosis factor binding protein fusion; ovary follicle cell maturation
    induction recombinant
TΤ
    Animal cell line
        (CHO, expression host; recombinant fusion proteins comprising
       ligand-binding receptor fragment linked with hormone subunit,
       heterodimer formation, and pharmaceutical uses)
ΙT
    Animal cell line
        (COS-7, expression host; recombinant fusion proteins comprising
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ligand-binding receptor fragment linked with hormone subunit,

heterodimer formation, and pharmaceutical uses) TT Plasmid vectors (D.alpha.-TBP190hCG.alpha.; recombinant fusion proteins comprising ligand-binding receptor fragment linked with hormone subunit, heterodimer formation, and pharmaceutical uses) ΙT Plasmid vectors (D.alpha.-TBP190hCG.beta.; recombinant fusion proteins comprising ligand-binding receptor fragment linked with hormone subunit, heterodimer formation, and pharmaceutical uses) ΙT Proteins, specific or class RL: BPN (Biosynthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (TBP1 (tumor necrosis factor binding protein 1), fusion products, with hormone subunit; recombinant fusion proteins comprising ligand-binding receptor fragment linked with hormone subunit, heterodimer formation, and pharmaceutical uses) TΤ Proteins, specific or class RL: BPN (Biosynthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (TBP2 (tumor necrosis factor binding protein 2), fusion products, with hormone subunit; recombinant fusion proteins comprising ligand-binding receptor fragment linked with hormone subunit, heterodimer formation, and pharmaceutical uses) ΙT Chimeric gene Chimeric gene RL: BPR (Biological process); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses) (animal; recombinant fusion proteins comprising ligand-binding receptor fragment linked with hormone subunit, heterodimer formation, and pharmaceutical uses) ΙT Gene, animal Gene, animal RL: BPR (Biological process); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses) (chimeric; recombinant fusion proteins comprising ligand-binding receptor fragment linked with hormone subunit, heterodimer formation, and pharmaceutical uses) ΙT Animal cell (expression host; recombinant fusion proteins comprising ligand-binding receptor fragment linked with hormone subunit, heterodimer formation, and pharmaceutical uses) TΤ Ovary (follicle cell, maturation induction; recombinant fusion proteins comprising ligand-binding receptor fragment linked with hormone subunit, heterodimer formation, and pharmaceutical uses) TT Interleukin 6 RL: BPN (Biosynthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (fusion products, with hormone receptor; recombinant fusion proteins comprising ligand-binding receptor fragment linked with hormone subunit, heterodimer formation, and pharmaceutical uses) Ligands TΤ Receptors RL: BPN (Biosynthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (fusion products, with hormone subunit; recombinant fusion proteins comprising ligand-binding receptor fragment linked with hormone subunit, heterodimer formation, and pharmaceutical uses) ΙT Antibodies FSH receptors Gonadotropin receptors RL: BPN (Biosynthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(fusion products; recombinant fusion proteins comprising ligand-binding receptor fragment linked with hormone subunit, heterodimer formation, and pharmaceutical uses)

IT Molecular association

(in heterodimer formation; recombinant fusion proteins comprising ligand-binding receptor fragment linked with hormone subunit, heterodimer formation, and pharmaceutical uses)

IT Peptides, biological studies

RL: BPN (Biosynthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(linker, contg. enzyme cleavage site; recombinant fusion proteins comprising ligand-binding receptor fragment linked with hormone subunit, heterodimer formation, and pharmaceutical uses)

IT Tumor necrosis factor receptors

RL: BAC (Biological activity or effector, except adverse); BPN (Biosynthetic preparation); BSU (Biological study, unclassified); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(p55, fusion products, with human chorionic gonadotropin; recombinant fusion proteins comprising ligand-binding receptor fragment linked with hormone subunit, heterodimer formation, and pharmaceutical uses)

IT Plasmid vectors

(pSVL-hTBP1.hCG.alpha.; recombinant fusion proteins comprising ligand-binding receptor fragment linked with hormone subunit, heterodimer formation, and pharmaceutical uses)

IT Plasmid vectors

(pSVL-hTBPhCG.beta.; recombinant fusion proteins comprising ligand-binding receptor fragment linked with hormone subunit, heterodimer formation, and pharmaceutical uses)

IT Enzymes, biological studies

RL: BSU (Biological study, unclassified); BIOL (Biological study) (peptide linker contg. enzyme cleavage site; recombinant fusion proteins comprising ligand-binding receptor fragment linked with hormone subunit, heterodimer formation, and pharmaceutical uses)

IT Ovary

(peptide linker contg. ovary enzyme cleavage site; recombinant fusion proteins comprising ligand-binding receptor fragment linked with hormone subunit, heterodimer formation, and pharmaceutical uses)

IT Tumor necrosis factors

RL: ADV (Adverse effect, including toxicity); BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)

(protection against cytotoxicity of; recombinant fusion proteins comprising ligand-binding receptor fragment linked with hormone subunit, heterodimer formation, and pharmaceutical uses)

IT DNA sequences

Drugs

Genetic vectors

Molecular cloning

Plasmid vectors

Protein sequences

(recombinant fusion proteins comprising ligand-binding receptor fragment linked with hormone subunit, heterodimer formation, and pharmaceutical uses)

IT Fusion proteins (chimeric proteins)

RL: BPN (Biosynthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(recombinant fusion proteins comprising ligand-binding receptor fragment linked with hormone subunit, heterodimer formation, and pharmaceutical uses)

IT Hormones, animal, biological studies

RL: BPN (Biosynthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

ΙT

ΙT

TΤ

ΙT

ΙT

ΙT

ΙT

ΙT

ΤТ

(subunit, fusion products; recombinant fusion proteins comprising ligand-binding receptor fragment linked with hormone subunit, heterodimer formation, and pharmaceutical uses) Interferon receptors RL: BPN (Biosynthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (.alpha.-interferon, fusion products; recombinant fusion proteins comprising ligand-binding receptor fragment linked with hormone subunit, heterodimer formation, and pharmaceutical uses) Interferons RL: BPN (Biosynthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (.beta., fusion products, with hormone receptor; recombinant fusion proteins comprising ligand-binding receptor fragment linked with hormone subunit, heterodimer formation, and pharmaceutical uses) Interferon receptors RL: BPN (Biosynthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (.beta., fusion products; recombinant fusion proteins comprising ligand-binding receptor fragment linked with hormone subunit, heterodimer formation, and pharmaceutical uses) Interferon receptors RL: BPN (Biosynthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (.gamma.-interferon, fusion products; recombinant fusion proteins comprising ligand-binding receptor fragment linked with hormone subunit, heterodimer formation, and pharmaceutical uses) 195460-72-1P **195460-74-3P** 195460-68-5P **195460-70-9P** RL: BAC (Biological activity or effector, except adverse); BPN (Biosynthetic preparation); BSU (Biological study, unclassified); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (amino acid sequence; recombinant fusion proteins comprising ligand-binding receptor fragment linked with hormone subunit, heterodimer formation, and pharmaceutical uses) 195460-67-4 195460-69-6 195460-71-0 195460-73-2 RL: BPR (Biological process); BSU (Biological study, unclassified); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses) (nucleotide sequence; recombinant fusion proteins comprising ligand-binding receptor fragment linked with hormone subunit, heterodimer formation, and pharmaceutical uses) 9002-04-4, Thrombin RL: BSU (Biological study, unclassified); BIOL (Biological study) (peptide linker contg. thrombin cleavage site; recombinant fusion proteins comprising ligand-binding receptor fragment linked with hormone subunit, heterodimer formation, and pharmaceutical uses) 9002-61-3DP, Human chorionic gonadotropin, subunit, fusion products RL: BAC (Biological activity or effector, except adverse); BPN (Biosynthetic preparation); BSU (Biological study, unclassified); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (recombinant fusion proteins comprising ligand-binding receptor fragment linked with hormone subunit, heterodimer formation, and pharmaceutical uses) 9002-67-9DP, Luteinizing hormone, subunit, fusion products 9002-68-0DP, Follicle stimulating hormone, subunit, fusion products 9002-71-5DP, Tsh hormone, subunit, fusion products 9014-42-0DP, Thrombopoietin, fusion products, with hormone receptor 57285-09-3DP, Inhibin, subunit, fusion products

RL: BPN (Biosynthetic preparation); THU (Therapeutic use); BIOL

(recombinant fusion proteins comprising ligand-binding receptor

(Biological study); PREP (Preparation); USES (Uses)

fragment linked with hormone subunit, heterodimer formation, and pharmaceutical uses)

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L26 ANSWER 8 OF 11 HCAPLUS COPYRIGHT 2002 ACS
AN
     1992:208630 HCAPLUS
 DN
     116:208630
ΤI
     Analogs of glycoprotein hormones having altered immunological
     characteristics, efficacy and/or receptor specificity
 ΤN
     Campbell, Robert K.; Moyle, William R.
 PΑ
     University of Medicine and Dentistry of New Jersey, USA
SO
     PCT Int. Appl., 93 pp.
     CODEN: PIXXD2
DT
     Patent
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LA
     ICM A61K037-38
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     2-4 (Mammalian Hormones)
     Section cross-reference(s): 3
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 PRAI US 1990-520703
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     WO 1991-US3162
                            19910507 <--
     Chimeric chorionic gonadotropin (CG) heterodimeric polypeptides are
AB
     provided which have different properties compared to native human CG
      (hCG). Certain heterodimeric polypeptides bind to LH (LH) and FSH
     receptors and stimulate steroidogenesis in testicular and ovarian cells.
     Other heterodimeric polypeptides bind to LH receptors but have lower
     efficacy than hCG in stimulation of steroidogenesis in testicular and
     ovarian cells. Prodn. of the chimeric analogs by recombinant techniques
     is described, and sequences of chimeras are included. The steroidogenesis
     potency of the analogs was strongly related to receptor binding activity.
     One analog had reduced efficacy, relative to hCG, for LH receptor-mediated
     cAMP accumulation; the analog also inhibited the ability of hCG to
     stimulate hCG-induced cAMP accumulation.
ST
     glycoprotein chimeric hormone analog; human chorionic gonadotropin
     chimeric analog; steroidogenesis chorionic gonadotropin analog; cyclic AMP
     chorionic gonadotropin analog; cloning chorionic gonadotropin analog cDNA
 TΤ
     Gene, animal
     RL: BIOL (Biological study)
         (cDNA, for chimeric chorionic gonadotropin .alpha. and .beta. subunits
        of human, expression in mammalian cells of)
 IT
     Cattle
     Fish
     Horse
     Sheep
         (chimeric heterodimeric glycoprotein hormone with sequences of human
IT
     Deoxyribonucleic acid sequences
         (of chimeric chorionic gonadotropin .alpha. and .beta. subunit cDNAs of
        human)
·IT
     Molecular cloning
         (of chimeric chorionic gonadotropin .alpha. and .beta. subunit cDNAs of
        human, in mammalian cells)
ΙT
     Protein sequences
         (of chimeric chorionic gonadotropin .alpha. and .beta. subunits of
```

human)

IT Plasmid and Episome

(pBMT2X-hCG-alpha, chimeric human chorionic gonadotropin .beta. subunit cDNA on, expression in mammalian cells of)

IT Plasmid and Episome

(pBMT2X-hCG-beta, for chimeric human chorionic gonadotropin .beta. subunit cDNA on, expression in mammalian cells of)

IT Plasmid and Episome

(pBNT2X-F8, chimeric human chorionic gonadotropin .beta. subunit cDNA on, expression in mammalian cells of)

IT Plasmid and Episome

(pCM-hCG-beta, chimeric human chorionic gonadotropin .beta. subunit cDNA on, expression in mammalian cells of)

IT Plasmid and Episome

(pCM-beta-J2, chimeric human chorionic gonadotropin .beta. subunit cDNA on, expression in mammalian cells of)

IT Plasmid and Episome

(pKBM-hCG-alpha, human chorionic gonadotropin .alpha. subunit cDNA on, chimeric .alpha. subunits manuf. in relation to)

IT Plasmid and Episome

(pKBM-hCG-beta, human chorionic gonadotropin .beta. subunit cDNA on, chimeric .beta. subunits manuf. in relation to)

IT Plasmid and Episome

(pKBM-hCG-beta', human chorionic gonadotropin .beta. subunit cDNA on, chimeric .beta. subunits manuf. in relation to)

IT Plasmid and Episome

(pSVL-B11, chimeric human chorionic gonadotropin .beta. subunit cDNA on, expression in mammalian cells of)

IT Plasmid and Episome

(pSVL-B9, chimeric human chorionic gonadotropin .beta. subunit cDNA on, expression in mammalian cells of)

IT Plasmid and Episome

(pSVL-F8, chimeric human chorionic gonadotropin .beta. subunit cDNA on, expression in mammalian cells of)

IT Plasmid and Episome

(pSVL-H3, chimeric human chorionic gonadotropin .alpha. subunit cDNA on, expression in mammalian cells of)

IT Plasmid and Episome

(pSVL-H6, chimeric human chorionic gonadotropin .alpha. subunit cDNA on, expression in mammalian cells of)

IT Plasmid and Episome

(pSVL-hCG-alpha, human chorionic gonadotropin .alpha. subunit cDNA on, chimeric .alpha. subunits manuf. in relation to)

IT Plasmid and Episome

(pSVL-hCG-beta, human chorionic gonadotropin .beta. subunit cDNA on, chimeric .beta. subunits manuf. in relation to)

IT Animal cell line

(C-127, expression in, of chimeric human chorionic gonadotropin .alpha. and .beta. subunit cDNAs)

IT Animal cell line

(COS, expression in, of chimeric human chorionic gonadotropin .alpha. and .beta. subunit cDNAs)

IT Animal cell line

(COS-7, expression in, of chimeric human chorionic gonadotropin .alpha. and .beta. subunit cDNAs)

IT Deoxyribonucleic acids

RL: BIOL (Biological study)

(complementary, for chimeric chorionic gonadotropin .alpha. and .beta. subunits of human, expression in mammalian cells of)

IT Glycoproteins, specific or class

RL: BIOL (Biological study)

(spike, G, fusion protein with human chorionic gonadotropin .beta.-chain fragment of, of vesicular stomatitis virus, prodn. in

```
COS-7 cells of)
TΤ
     Virus, animal
        (vesicular stomatitis, G protein of, fusion products with human
        chorionic gonadotropin .beta.-chain fragment, prodn. in COS-7 cells of)
                   140933-24-0
                                 140933-25-1 140933-26-2 140933-28-4
ΤТ
     140933-23-9
     140933-29-5
                   140933-31-9
     RL: PRP (Properties)
        (amino acid sequence of, complete, and monoclonal antibody binding
        activity of)
     140933-32-0D, fusion products with vesicular stomatitis virus G protein
ΤТ
     RL: PRP (Properties)
        (amino acid sequence of, complete, and monoclonal antibody binding to
        cell expressing)
     140933-27-3
ΙT
     RL: PRP (Properties)
        (amino acid sequence of, complete, chimeric chorionic gonadotropin
        heterodimer manuf. in relation to)
     140933-02-4
                   140933-04-6
                                 140933-06-8
IT
                                               140933-07-9
                                                              140933-08-0
                                 140933-11-5
     140933-09-1
                   140933-10-4
                                               140933-12-6
     RL: PRP (Properties)
        (amino acid sequence of, complete, receptor binding of)
TΨ
     140933-21-7P
                    140933-22-8P
                                   140933-30-8P
     RL: PRP (Properties); PREP (Preparation)
        (amino acid sequence of, complete, recombinant prodn. and altered
        activity of)
     140933-03-5P
TΤ
                    140933-13-7P
     RL: PRP (Properties); PREP (Preparation)
        (amino acid sequence of, complete, recombinant prodn. and receptor
        binding of)
ΤТ
     140933-85-3, Deoxyribonucleic acid (ox prechorionic gonadotropin
     .alpha.-subunit-specifying plus 5'- and 3'-flanking region fragment)
     RL: PRP (Properties)
        (nucleotide sequence of, chimeric chorionic gonadotropin heterodimer
        manuf. in relation to)
ΙT
     140933-84-2, Deoxyribonucleic acid (ox prechorionic gonadotropin
     .alpha.-subunit-specifying)
     RL: PRP (Properties)
        (nucleotide sequence of, complete, chimeric chorionic gonadotropin
        heterodimer manuf. in relation to)
     140933-68-2, Deoxyribonucleic acid (human prechorionic gonadotropin
IT
     .beta.-subunit-specifying)
     RL: PRP (Properties)
        (nucleotide sequence of, complete, chimeric .beta.-chain cDNA prepn.
        using)
                                 140933-67-1
TΤ
     140933-33-1
                   140933-66-0
     RL: PRP (Properties)
        (nucleotide sequence of, complete, recombinant heterodimeric chorionic
        gonadotropin manuf. in relation to)
     76050-53-8
TΤ
                  79030-14-1
     RL: BIOL (Biological study)
        (recombinant chimeric chorionic gonadotropin heterodimers prepn. using)
     9002-61-3D, Chorionic gonadotropin, .beta. subunit fragment, fusion
ΙT
     products with FSH or TSH fragment
                                         9002-68-0D, Follicle-stimulating
     hormone, .beta. subunit fragment, fusion products with chorionic
     gonadotropin .beta. subunit fragment
                                            9002-71-5D, Thyroid-stimulating
     hormone, .beta. subunit fragment, fusion products with chorionic
     gonadotropin .beta. subunit fragment
     RL: BIOL (Biological study)
        (recombinant, altered receptor binding of)
L26 ANSWER 9 OF 11 HCAPLUS COPYRIGHT 2002 ACS
     1984:97541 HCAPLUS
ΑN
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DN

100:97541

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TI
     Evolution of the genes for the .beta. subunits of human chorionic
     gonadotropin and luteinizing hormone
     Talmadge, Karen; Vamvakopoulos, Nikos C.; Fiddes, John C.
ΑU
     Cold Spring Harbor Lab., Cold Spring Harbor, NY, 11724, USA Nature (London) (1984), 307(5946), 37-40
CS
SO
     CODEN: NATUAS; ISSN: 0028-0836
DT
     Journal
     English
LA
CC
     3-3 (Biochemical Genetics)
     Section cross-reference(s): 2, 13
AΒ
     Nucleotide sequence comparisons of the single gene for the human LH
     [9002-67-9] gene .beta.-subunit with 2 of the 7 genes for the human
     chorionic gonadotropin [9002-61-3] .beta.-subunit suggest that the .beta.
     human chorionic gonadotropin genes have evolved from an ancestral .beta.
     LH gene by a series of selected changes with very little neutral drift.
     Moreover, the 24-amino acid C-terminal extension of the human chorionic
     gonadotropin .beta.-subunit appears to have arisen by a single base
     deletion that incorporated the 3'-untranslated region of the ancestral
     .beta. LH gene into the coding region.
ST
     chorionic gonadotropin LH gene human evolution
ΙT
     Gene and Genetic element, animal
     RL: PROC (Process)
        (for chorionic gonadotropin and LH .beta.-subunits, of human, structure
        and evolution of)
ΙT
     Protein sequences
        (of LH .beta.-subunit, of human, complete)
TT
     Evolution
        (of chorionic gonadotropin and LH .beta.-subunit genes, of human)
IT
     Protein sequences
        (of chorionic gonadotropin .beta.-subunit precursor, of human,
        complete)
     Protein sequences
IT
        (of chorionic gonadotropin .beta.-subunit, of human, complete)
ΙT
     Deoxyribonucleic acid sequences
        (LH .beta.-subunit-specifying, of human, complete)
     Deoxyribonucleic acid sequences
ΙT
        (chorionic gonadotropin .beta.-subunit-specifying, of human, complete)
                  56832-34-9 76050-53-8 87971-06-0 87971-07-1
TT
     53664-53-2
     89072-90-2
     RL: PRP (Properties)
        (amino acid sequence of)
     9002-61-3
                 9002-67-9
IT
     RL: PRP (Properties)
        (gene for .beta.-subunit of, of human, structure and evolution of)
     89072-67-3
                  89072-68-4
                               89072-69-5
ΙT
     RL: PRP (Properties); BIOL (Biological study)
        (nucleotide sequence of)
     ANSWER 10 OF 11 HCAPLUS COPYRIGHT 2002 ACS
L26
ΑN
     1983:607297 HCAPLUS
DN
     99:207297
     The .beta. subunit of human chorionic gonadotropin is encoded by multiple
ΤI
ΑU
     Policastro, Paul; Ovitt, Catherine E.; Hoshina, Makoto; Fukuoka, Hideoki;
     Boothby, Mark R.; Boime, Irving
CS
     Sch. Med., Washington Univ., St. Louis, MO, 63110, USA
     J. Biol. Chem. (1983), 258(19), 11492-9
SO
     CODEN: JBCHA3; ISSN: 0021-9258
\mathsf{DT}
     Journal
LA
     English
CC
     3-3 (Biochemical Genetics)
     Section cross-reference(s): 2, 13
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Two recombinant phage clones bearing sequences corresponding to the .beta.

AB

subunit of human chorionic gonadotropin (hCG.beta.) [9002-61-3] were isolated from a human genomic library. The .beta. sequences were mapped by blot hybridization of restriction digests of these phage DNAs and the nonoverlapping inserts were subcloned in plasmid pBR322 and sequenced. The nucleotide-sequencing data show that the hCG.beta. subunit is encoded by .gtoreq.3 nonallelic genes. Moreover, restriction analyses of human placental DNA indicated that these genes may be linked in a single cluster with 4 other hCG.beta.-like genes. The sequenced genes all differ in their 5' flanking regions, and none of them is completely homologous in sequence to either of the 2 hCG.beta. clones used here. In the translated region of 1 of these genes, 3 base substitutions result in 2 changes from the reported amino acid sequence. In the family of .beta.-contg. glycoprotein hormones, the hCG.beta. subunits is unique in that it contains an extension of 29 amino acids at its C-terminus. The DNA sequence corresponding to this region in the sequenced genes is part of a larger exon. The C-terminal extension does not result from splicing of the primary RNA transcript.

ST human chorionic gonadotropin beta subunit; gene human chorionic gonadotropin subunit; sequence human chorionic gonadotropin subunit

IT Gene and Genetic element, animal

RL: BIOL (Biological study)

(for chorionic gonadotropin .beta. subunit, of human, multiple)

IT Protein sequences

(of chorionic gonadotropin .beta. subunit precursor, of human multiple clones, complete)

IT Protein sequences

(of chorionic gonadotropin .beta. subunit, of human multiple clones, complete)

IT Deoxyribonucleic acid sequences

(chorionic gonadotropin .beta.-subunit-specifying, of human genomic multiple clones, complete)

IT 56832-34-9 **76050-53-8** 87971-06-0 87971-07-1

RL: PRP (Properties)

(amino acid sequence of)

IT 76012-21-0 87970-97-6

RL: PRP (Properties); BIOL (Biological study)
 (nucleotide sequence of)

IT 9002-61-3

RL: PRP (Properties)

(.beta. subunit of, of human, multiple genes for)

- L26 ANSWER 11 OF 11 HCAPLUS COPYRIGHT 2002 ACS
- AN 1981:11741 HCAPLUS
- DN 94:11741
- TI The cDNA for the .beta.-subunit of human chorionic gonadotropin suggests evolution of a gene by readthrough into the 3'-untranslated region
- AU Fiddes, John C.; Goodman, Howard M.
- CS Howard Hughes Med. Inst. Lab., Univ. California, San Francisco, CA, 94143, USA
- SO Nature (London) (1980), 286(5774), 684-7 CODEN: NATUAS; ISSN: 0028-0836
- DT Journal
- LA English
- CC 6-2 (General Biochemistry)
- AB A 579-base pair approx. full-length complementary DNA (cDNA) coding for the 145-amino acid long .beta.-subunit of human chorionic gonadotropin (I) was cloned in the plasmid vector pBR322 and its complete nucleotide sequence detd. A hydrophobic presequence of 20 amino acids was identified from the nucleotide sequence. The amino acid sequence of the .beta.-subunit of I contained a C-terminal extension of .apprx.30 amino acids which has no homologous counterpart in LH, FSH, and TSH, although the amino acid sequence of the .beta.-subunit is related to those of the .beta.-subunits of LH, FSH, and TSH. Anal. of the nucleotide sequence of

ST

TT

IT

IT

IT

ΙT

TΤ

ΙT

ΙT

TΤ

TT

ΑN DN

TΙ

ΙN

PΑ

SO

DT

LA

IC

CC

2-7 (Mammalian Hormones)

.beta.-subunit of I cDNA suggested that this extension may have arisen by the loss of the termination codon of an ancestral .beta.-like gene so that most of what was previously the 3'-untranslated region now codes for protein. The .beta.-subunit of I terminated with the codon UAA located 16 bases before the poly(A) in the sequence AAUAAA. This sequence may be a recognition signal involved in either polyadenylation or processing and therefore may have a dual role in this gene, serving both a coding and regulatory function. chorionic gonadotropin beta subunit gene; evolution gene chorionic gonadotropin subunit; nucleotide sequence chorionic gonadotropin gene; complementary DNA chorionic gonadotropin sequence Peptides, properties RL: PRP (Properties) (amino acid sequence of, of chorionic gonadotropin (human .beta.-subunit precursor reduced)) Gene RL: BIOL (Biological study) (for chorionic gonadotropin .beta.-subunit of human, evolution of, complementary DNA nucleotide sequence in relation to) Molecular structure, natural product (of DNA (human chorionic gonadotropin .beta.-subunit precursor complementary)) Molecular structure, natural product (of chorionic gonadotropin (human .beta.-subunit precursor reduced)) Evolution (of chorionic gonadotropin .beta.-subunit gene, nucleotide sequence of complementary DNA in relation to) Nucleotides, properties RL: PRP (Properties) (sequence of, of complementary DNA for .beta.-subunit of human chorionic gonadotropin) Deoxyribonucleic acids RL: BIOL (Biological study) (complementary, for chorionic gonadotropin .beta.-subunit of human, nucleotide sequence of, gene evolution in relation to) 76050-53-8 RL: PRP (Properties) (amino acid sequence of) 56832-34-9 RL: BIOL (Biological study) (complementary DNA for .beta.-subunit of, nucleotide sequence of, gene evolution in relation to) 76012-21-0 RL: PRP (Properties) (nucleotide sequence of) => d all 124 L24 ANSWER 1 OF 1 HCAPLUS COPYRIGHT 2002 ACS 1999:223048 HCAPLUS 130:247459 Mutants of thyroid stimulating hormone subunits with improved bioactivity and stability Weintraub, Bruce D.; Szkudlinski, Mariusz W. University of Maryland, Baltimore, USA PCT Int. Appl., 44 pp. CODEN: PIXXD2 Patent English ICM C12N015-16 ICS C07K014-59; A61K038-24; G01N033-68

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FAN.CNT 1
                      KIND DATE
     PATENT NO.
                                           APPLICATION NO.
                                                             DATE
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                       A2
                            19990401
                                           WO 1998-US19772
                                                            19980922
                      A3
     WO 9915665
                            19990520
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             RU, TJ,
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                     GA, GN, GW, ML, MR, NE, SN, TD, TG
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             IE, FI
PRAI US 1997-939472
                       Α2
                            19970922
     WO 1998-US19772
                       W
                            19980922
                       W
     WO 1999-US5908
                            19990319
                                      <--
AΒ
     The present invention is based upon the discovery that mutant .alpha.
     subunits and mutant .beta. subunits each comprising amino acid
     substitutions relative to the wild type can be produced and assembled to
     form a mutant TSH heterodimer or TSH analog that possesses higher
     bioactivity in vitro and longer half life in vivo. A preferred mutant
     .alpha. subunit (to be used in conjunction with a modification to increase
     the serum half-life of the TSH heterodimer having the mutant .alpha.
     subunit) comprises four mutations: Q13K, E14K, P16K, and Q20K; a preferred
     mutant .beta. subunit comprises three mutations: I58R, E63R, and L69R.
     Multiple mutations within a subunit and modifications to increase the
     half-life of the TSH heterodimer (i.e., .beta.-subunit fusion with the
     C-terminal extension peptide of human chorionic gonadotropin and/or a
     .beta. subunit-.alpha. subunit fusion) can act synergistically to achieve
     bioactivity that is greater than the sum of the increase of the mutations
     and the long acting modifications. Accordingly, the present invention
     provides methods for using mutant TSH heterodimers, TSH analogs,
     fragments, and derivs. thereof for treating or preventing diseases of the
     thyroid, in particular thyroid cancer. The invention also relates to
     methods of diagnosis, prognosis and monitoring for thyroid-related
     functions. Pharmaceutical and diagnostic compns., methods of using mutant
     TSH heterodimers and TSH analogs with utility for treatment and prevention
     of metabolic and reproductive diseases are also provided.
ST
     TSH mutagenesis bioactivity stability
ΙT
     Diagnosis
        (cancer; mutants of human TSH subunits with improved bioactivity and
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stability)

Antibodies

ΙT

RL: ANT (Analyte); ANST (Analytical study) (diagnosis of antibodies against TSH receptor in Graves' disease; mutants of human TSH subunits with improved bioactivity and stability) IT Thyrotropin receptors RL: BSU (Biological study, unclassified); BIOL (Biological study) (diagnosis of antibodies against TSH receptor in Graves' disease; mutants of human TSH subunits with improved bioactivity and stability) Graves' disease IT(diagnosis of; mutants of human TSH subunits with improved bioactivity and stability) Neoplasm TT (diagnosis; mutants of human TSH subunits with improved bioactivity and stability) Test kits TT (for diagnosis of Graves' disease; mutants of human TSH subunits with improved bioactivity and stability) IT Thyroid gland, neoplasm Thyroid gland, neoplasm (inhibitors; mutants of human TSH subunits with improved bioactivity and stability) IT Diagnosis (mol.; mutants of human TSH subunits with improved bioactivity and stability) Mutagenesis TT Protein engineering (mutants of human TSH subunits with improved bioactivity and stability) TΤ Protein sequences (of mutants of human TSH subunits with improved bioactivity and stability) Antitumor agents IT Antitumor agents (thyroid; mutants of human TSH subunits with improved bioactivity and stability) TΤ Hypothyroidism (treatment of; mutants of human TSH subunits with improved bioactivity and stability) ΙT 9002-71-5DP, Thyroid stimulating hormone, mutants 56832-30-5DP, mutants 64365-92-0DP, Thyrotropin (human .beta.-subunit protein moiety reduced), 221650-43-7P 221650-44-8P 221650-45-9P 221650-46-0P mutants 221650-47-1P 221650-48-2P 221650-49-3P 221650-50-6P 221650-51-7P 221650-52-8P 221650-53-9P RL: BAC (Biological activity or effector, except adverse); BPN (Biosynthetic preparation); BSU (Biological study, unclassified); PRP

(Properties); BIOL (Biological study); PREP (Preparation)

(mutants of human TSH subunits with improved bioactivity and stability)